SPECIFICATION AMENDMENTS

Please amend the ABSTRACT as follows:

ABSTRACT

The present invention provides chimeric GPCRs that display novel downstream signaling properties Edg receptors, useful as components of an assay system. The present invention also provide methods of screening for compounds that bind a chimeric Edg receptor, for example, by contacting a chimeric Edg receptor with a compound and detecting a change in chimeric Edg receptor-mediated activity such as calcium mobilization.



SPECIFICATION AMENDMENTS

Please amend the specification as follows:

Amend the paragraph beginning on page 7, line 18, as indicated by the underlined portion, and amend the subsequent table by adding the words "<u>Table 1</u>" and enlarging the font size of the lettering used in the table, as shown below:

The amino acid notations used herein for the twenty genetically encoded L-amino acids are conventional and are as shown in Table 1. follows:

Table 1

Amino Acid	One-Letter Abbreviation	Three Letter Abbreviation
Alanine	Α	Ala
Arginine	R	Arg
Asparagine	N	Asn
Aspartic acid	D	Asp
Cysteine	C	Cys
Glutamine	Q E	Gln
Glutamic acid	Ē	Glu
Glycine	. G	Gly
Histidine	H	His
Isoleucine	I	Ile
Leucine	L	Leu
Lysine	K	Lys
Methionine	M	Met
Phenylalanine	F	Phe
Proline	P	Pro
Serine	S	Ser
Threonine	T	Thr
Tryptophan	W	Trp
Tyrosine	Y	Tyr
Valine	V	Val



Table 2

Amino Acid Sequences of Edg Receptors

Human Edg 1 (SEQ ID NO: 1)

Genbank Accession No. AF233365 MGPTSVPLVKAHRSSVSDYVNYDIIVRHYNYTGKLNISAD
KENSIKLTSVVFILICCFIILENIFVLLTIWKTKKFHRPM
YYFIGNLALSDLLAGVAYTANLLLSGATTYKLTPAQWFLR
EGSMFVALSASVFSLLAIAIERYITMLKMKLHNGSNNFRL
FLLISACWVISLILGGLPIMGWNCISALSSCSTVLPLYHK
HYILFCTTVFTLLLLSIVILYCRIYSLVRTRSRRLTFRKN
ISKASRSSEKSLALLKTVIIVLSVFIACWAPLFILLLLDV
GCKVKTCDILFRAEYFLVLAVLNSGTNPIIYTLTNKEMRR
AFIRIMSCCKCPSGDSAGKFKRPIIAGMEFSRSKSDNSSH
PQKDEGDNPETIMSSGNVNSSS

Human Edg2 (SEQ ID NO:30)

Genbank Accession No. U78192 MAAISTSIPVISQPQFTAMNEPQCFYNESIAFFYNRSGKH
LATEWNTVSKLVMGLGITVCIFIMLANLLVMVAIYVNRRF
HFPIYYLMANLAAADFFAGLAYFYLMFNTGPNTRRLTVST
WLLRQGLIDTSLTASVANLLAIAIERHITVFRMQLHTRMS
NRRVVVVIVVIWTMAIVMGAIPSVGWNCICDIENCSNMAP
LYSDSYLVFWAIFNLVTFVVMVVLYAHIFGYVRQRTMRMS
RHSSGPRNRDTMMSLLKTVVIVLGAFIICWTPGLVLLLL
DVCCPQCDVLAYEKFFLLLAEFNSAMNPIIYSYRDKEMSA
TFRQILCCQRSENPTGPTEGSDRSASSLNHTILAGVHSND
HSVV

Human Edg 3 (SEQ ID NO:2)

Genbank Accession No. X83864 MATALPPRLQPVRGNETLREHYQYVGKLAGRLKEASEGST
LTTVLFLVICSFIVLENLMVLIAIWKNNKFHNRMYFFIGN
LALCDLLAGIAYKVNILMSGKKTFSLSPTVWFLREGSMFV
ALGASTCSLLAIAIERHLTMIKMRPYDANKRHRVFLLIGM
CWLIAFTLGALPILGWNCLHNLPDCSTILPLYSKKYIAFC
ISIFTAILVTIVILYARIYFLVKSSSRKVANHNNSERSMA
LLRTVVIVVSVFIACWSPLFILFLIDVACRVQACPILFKA
QWFIVLAVLNSAMNPVIYTLASKEMRRAFFRLVCNCLVRG
RGARASPIQPALDPSRSKSSSSNNSSHSPKVKEDLPHTDP
SSCIMDKNAALQNGIFCN

B3

Human Edg4 (SEQ ID NO:31)

Genbank Accession No. AF233092

MVIMGQCYYNETIGFFYNNSGKELSSHWRPKDVVVVALGL TVSVLVLLTNLLVIAAIASNRRFHQPIYYLLGNLAAADLF AGVAYLFLMFHTGPRTARLSLEGWFLROGLLDTSLTASVA TLLAIAV*ERHRSVMAVQLHSRLPRGR*VVMLIVGVWVAALG LGLLPAHSWHCLCALDRCSRMAPLLSRSYLAVWALSSLLV FLLMVAVYTRIFFYV*RRRVORMAEHVSCHPRYRETTLSLV* KTVVIILGAFVVCWTPGQVVLLLDGLGCESCNVLAVEKYF ${ t LLLAEANSLVNAAVYSC} RDAEMRRTFRRLLCCACLRQSTR$ *ESVHYTSSAQGGASTRIMLPENGHPLMDSTL*

(SEQ ID NO:32)

Genbank Accession No. AF011466

Human Edg 4 mt MVIMGQCYYNETIGFFYNNSGKELSSHWRPKDVVVVALGL TVSVLVLLTNLLVIAAIASNRRFHQPIYYLLGNLAAADLF AGVAYLFLMFHTGPRTARLSLEGWFLROGLLDTSLTASVA TLLAIAV*ERHRSVMAVQLHSRLPRGR*VVMLIVGVWVAALG LGLLPAHSWHCLCALDRCSRMAPLLSRSYLAVWALSSLLV FLLMVAVYTRIFFYV*RRRVQRMAEHVSCHPRYRETTLSLV* KTVVIILGAFVVCWTPGQVVLLLDGLGCESCNVLAVEKYF LLLAEANSLVNAAVYSCRDAEMRRTFRRLLCCACLRQSTR ESVHYTSSAQGGASTRIMLPENGHPLMTPPFSYLELQRYA*ASNKSTAPDDLWVLLAQPNQQD*

Human Edg 5 (SEQ ID NO:33)

Genbank Accession No. AF034780

MGSLYSEYLNPNKVOEHYNYTKETLETOETTSROVASAFI VILCCAIVVENLLVLIAV*ARNSKFHS*AMYLFLGNLAASDL LAGVAFVANTLLSGSVTLRLTPVQWFAREGSASITLSASV FSLLAIAI*ERHVAIAKVKLYGSDKSCR*MLLLIGASWLISL VLGGLPILGWNCLGHLEACSTVLPLYAKHYVLCVVTIFSI ILLAIVALYVRIYCVV*RSSHADMAAPOTLALLK*TVTIVLG VFIVCWLPAFSILLLDYACPVHSCPILYKAHYFFAVSTLN SLLNPVIYTW*RSRDLRREVLRPLQCWRPGVGVQGRRRVGT* PGHHLLPLRSSSSLERGMHMPTSPTFLEGNTVV

Human Edg6 (SEQ ID NO:34)

Genbank Accession No. AJ000479

MNATGTPVAPESCQQLAAGGHSRLIVLHYNHSGRLAGRGG PEDGGLGALRGLSVAASCLVVLENLLVLAAI TSHMRSRRW VYYCLVNITLSDLLTGAAYLANVLLSGARTFRLAPAQWFL REGLLFTALAASTFSLLFTAG*ERFATMVRPVAESGATKTS* RVYGFIGLCWLLAALLGMLPLLGWNCLCAFDRCSSLLPLY SKRYILFCLVIFAGVLATIMGLYGAIFRLV*QASGQKAPRP AARRKARRLLK*TVLMILLAFLVCWGPLFGLLLADVFGSNL WAQEYLRGMDWILALAVLNSAVNPIIYSFRSREVCRAVLS FLCCGCLRLGMRGPGDCLARAVEAHSGASTTDSSLRPRDSFRGSRSLSFRMREPLSSISSVRSI



Human Edg7 (SEQ ID NO:35)

Genbank Accession No. AF127138 MNECHYDKHMDFFYNRSNTDTVDDWTGTKLVIVLCVGTFF
CLFIFFSNSLVIAAVIKNRKFHFPFYYLLANLAAADFFAG
IAYVFLMFNTGPVSKTLTVNRWFLRQGLLDSSLTASLTNL
LVIAVERHMSIMRMRVHSNLTKKRVTLLILLVWAIAIFMG
AVPTLGWNCLCNISACSSLAPIYSRSYLVFWTVSNLMAFL
IMVVVYLRIYVYVKRKTNVLSPHTSGSISRRRTPMKLMKT
VMTVLGAFVVCWTPGLVVLLLDGLNCRQCGVQHVKRWFLL
LALLNSVVNPIIYSYKDEDMYGTMKKMICCFSQENPERRP
SRIPSTVLSRSDTGSQYIEDSISQGAVCNKSTS

Human Edg 8 (SEQ ID NO:36)

Genbank Accession No. AF317676 MESGLLRPAPVSEVIVLHYNYTGKLRGARYQPGAGLRADA
VVCLAVCAFIVLENLAVLLVLGRHPRFHAPMFLLLGSLTL
SDLLAGAAYAANILLSGPLTLKLSPALWFAREGGVFVALT
ASVLSLLAIALERSLTMARRGPAPVSSRGRTLAMAAAAWG
VSLLLGLLPALGWNCLGRLDACSTVLPLYAKAYVLFCVLA
FVGILAAICALYARIYCQVRANARRLPARPGTAGTTSTRA
RRKPRSLALLRTLSVVLLAFVACWGPLFLLLLLDVACPAR
TCPVLLQADPFLGLAMANSLLNPIIYTLTNRDLRHALLRL
VCCGRHSCGRDPSGSQQSASAAEASGGLRRCLPPGLDGSF
SGSERSSPQRDGLDTSGSTGSPGAPTAARTLVSEPAAD

Amend Table 3, beginning on page 21, line 1, by changing the font size and spacing as indicated below (no additions or deletions are to be made to Table 3):

Table 3

Amino Acid Sequences of Chimeric Edg Receptors

Edg1/3(ct) (SEQ ID NO:3) MGPTSVPLVKAHRSSVSDYVNYDIIVRHYNYTGKLNISAD
KENSIKLTSVVFILICCFIILENIFVLLTIWKTKKFHRPM
YYFIGNLALSDLLAGVAYTANLLLSGATTYKLTPAQWFLR
EGSMFVALSASVFSLLAIAIERYITMLKMKLHNGSNNFRL
FLLISACWVISLILGGLPIMGWNCISALSSCSTVLPLYHK
HYILFCTTVFTLLLLSIVILYCRIYSLVRTRSRRLTFRKN
ISKASRSSEKSLALLKTVIIVLSVFIACWAPLFILLLLDV
GCKVKTCDILFRAEYFLVLAVLNSGTNPIIYTLTSKEMRR
AFFRLVCNCLVRGRGARASPIQPALDPSRSKSSSSNNSSH
SPKVKEDLPHTDPSSCIMDKNAALQNGIFCN



Edg1/3(i3ct) (SEQ ID NO:4) MGPTSVPLVKAHRSSVSDYVNYDIIVRHYNYTGKLNISAD
KENSIKLTSVVFILICCFIILENIFVLLTIWKTKKFHRPM
YYFIGNLALSDLLAGVAYTANLLLSGATTYKLTPAQWFLR
EGSMFVALSASVFSLLAIAIERYITMLKMKLHNGSNNFRL
FLLISACWVISLILGGLPIMGWNCISALSSCSTVLPLYHK
HYILFCTTVFTLLLLSIVILYCRIYSLVRSSSRKVANHNN
SERSMALLRTVIIVLSVFIACWAPLFILLLLDVGCKVKTC
DILFRAEYFLVLAVLNSGTNPIIYTLTSKEMRRAFFRLVC
NCLVRGRGARASPIQPALDPSRSKSSSSNNSSHSPKVKED
LPHTDPSSCIMDKNAALQNGIFCN

Edg1/3(i2i3ct) (SEQ ID NO:5)

MGPTSVPLVKAHRSSVSDYVNYDIIVRHYNYTGKLNISAD
KENSIKLTSVVFILICCFIILENIFVLLTIWKTKKFHRPM
YYFIGNLALSDLLAGVAYTANLLLSGATTYKLTPAQWFLR
EGSMFVALSASVFSLLAIAIERHLTMIKMRPYDANKRHRL
FLLISACWVISLILGGLPIMGWNCISALSSCSTVLPLYHK
HYILFCTTVFTLLLLSIVILYCRIYSLVRSSSRKVANHNN
SERSMALLRTVIIVLSVFIACWAPLFILLLLDVGCKVKTC
DILFRAEYFLVLAVLNSGTNPIIYTLTSKEMRRAFFRLVC
NCLVRGRGARASPIQPALDPSRSKSSSSNNSSHSPKVKED
LPHTDPSSCIMDKNAALQNGIFCN

Edg 5/3(i3ct) (SEQ ID NO:37)

MGSLYSEYLNPNKVQEHYNYTKETLETQETTSRQVASAFI
VILCCAIVVENLLVLIAVARNSKFHSAMYLFLGNLAASDL
LAGVAFVANTLLSGSVTLRLTPVQWFAREGSASITLSASV
FSLLAIAIERHVAIAKVKLYGSDKSCRMLLLIGASWLISL
VLGGLPILGWNCLGHLEACSTVLPLYAKHYVLCVVTIFSI
ILLAIVALYVRIYCVVKSSSRKVANHNNSERSMALLRTVT
IVLGVFIVCWLPAFSILLLDYACPVHSCPILYKAHYFFAV
STLNSLLNPVIYTWASKEMRRAFFRLVCNCLVRGRGARAS
PIQPALDPSRSKSSSSNNSSHSPKVKEDLPHTDPSSCIMD
KNAALQNGIFCN

Edg 8/4(ct) (SEQ ID NO:38) MESGLLRPAPVSEVIVLHYNYTGKLRGARYQPGAGLRADA
VVCLAVCAFIVLENLAVLLVLGRHPRFHAPMFLLLGSLTL
SDLLAGAAYAANILLSGPLTLKLSPALWFAREGGVFVALT
ASVLSLLAIALERSLTMARRGPAPVSSRGRTLAMAAAAWG
VSLLLGLLPALGWNCLGRLDACSTVLPLYAKAYVLFCVLA
FVGILAAICALYARIYCQVRANARRLPARPGTAGTTSTRA
RRKPRSLALLRTLSVVLLAFVACWGPLFLLLLLDVACPAR
TCPVLLQADPFLGLAMANSLLNPIIYTLRDAEMRRTFRRL
LCCACLRQSTRESVHYTSSAQGGASTRIMLPENGHPLMTP
PFSYLELQRYAASNKSTAPDDLWVLLAQPNQQD



Amend Table 4, beginning on page 35, line 13, by changing the font size and spacing as indicated below (no additions or deletions are to be made to Table 4):

<u>Table 4</u>

PCR Primers for Generating Chimeric Edg 1 Receptors

<u>Primer</u> Edg-1	Direction Positio	CCC/GCG/GTT/AAC/ATG/GGG/CCC/ACC/
(SEQ ID NO:6)		AGC/GTC
Edg-3 (SEQ ID NO:7)	rev 1137	CGC/GGA/TCC/TCA/GTT/GCA/GAA/GAT/CCC
E1/3 CTD (SEQ ID NO:8)	942	CAT/TTA/CAC/TCT/GAC/CAG/CAA/GGA/ GAT/GCG/GCG/G
E1/3 CTD (SEQ ID NO:9)	rev 942	CCG/CAT/CTC/CTT/GCT/GGT/CAG/AGT/ GTA/AAT/GAT/G
E1/3 i2 (SEQ ID NO:10)	402	GTC/TCC/TCG/CCA/TCG/CCA/TCG/AGC/GGC/ACT/TGA/C
E1/3 i2 (SEQ ID NO:11)	rev 402	GTC/AAG/TGC/CGC/TCG/ATG/GCG/ATG/GCG/AGG/AGA
E1/3 i2 (SEQ ID NO:12)	441	CGC/CAA/CAA/GAG/GCA/CCG/CCT/CTT/ CCT/GCT/AAT/C
E1/3 i2 (SEQ ID NO:13)	rev 441	GAT/TAG/CAG/GAA/GAG/GCG/GTG/CCT/ CTT/GTT/GGC/G
E1/3 i3 (SEQ ID NO:14)	684	CTA/CTC/CTT/GGT/CAG/GTC/CAG/CAG/ CCG/TAA/GGT/G
E1/3 i3 (SEQ ID NO:15)	rev 684	CAC/CTT/ACG/GCT/GCT/GGA/CCT/GAC/ CAA/GGA/GTA/G
E1/3 i3 (SEQ ID NO:16)	723	CAC/TGC/TGC/GGA/CCG/TGA/TTA/TCG/ TCC/TGA/GCG/TC
E1/3 i3 (SEQ ID NO:17)	rev 723	GAC/GCT/CAG/GAC/GAT/AAT/CAC/GGT/ CCG/CAG/CAG/TG

Amend Table 5, beginning on page 38, line 1, by changing the font size and spacing as indicated below (no additions or deletions are to be made to Table 5):

<u>Table 5</u>

PCR Primers for Generating Chimeric Edg 5 Receptors

Primer Edg-5 (SEQ ID NO:18)	Direction	Position 1	Sequence 5'-3' CCC/GCG/GTT/AAC/ATG/GGC/AGC/ TTG/TAC/TCG
Edg-3 (SEQ ID NO:19)	rev	1137	CGC/GGA/TCC/TCA/GTT/GCA/GAA/GAT/CCC
E5/3 (SEQ ID NO:20)		864	CGT/CAT/CTA/CAC/GTG/GGC/CAG/ CAA/GGA/GAT/GCG/G
E5/3 (SEQ ID NO:21)	rev	864	CCG/CAT/CTC/CTT/GCT/GGC/CCA/ CGT/GTA/GAT/GAC/G
E5/3 i3 (SEQ ID NO:22)		633	CAT/CTA/CTG/CGT/GGT/CAA/GTC/ CAG/CAG/CCG/TAA/G
E5/3 i3 (SEQ ID NO:23)	rev	633	CTT/ACG/GCT/GCT/GGA/CTT/GAC/CAC/GCA/GTA/GAT/G
E5/3 i3 (SEQ ID NO:24)		723	CAC/TGC/TGC/GGA/CCG/TGA/CCA/ TCG/TGC/TAG/GCG/TC
E1/3 i3 (SEQ ID NO:25)	rev	723	GAC/GCC/TAG/CAC/GAT/GGT/CAC/GGT/CCG/CAG/CAG/TG



Amend Table 6, beginning on page 39, line 8, by changing the font size and spacing as indicated below (no additions or deletions are to be made to Table 6):

Table 6 PCR Primers for Generating Chimeric Edg 8 Receptors

<u>Primer</u> Edg-8	Direction	Position 1	<u>Sequence 5'-3'</u> CCC/GCG/GTT/AAC/ATG/GAG/TCG/
(SEQ ID NO:26)			GGG/CTG/CTG
Edg-4-mut (SEQ ID NO:27)	rev	1149	CGC/GGA/TCC/TCA/GTC/CTG/TTG/ GTT/GGG
E8/4 (SEQ ID NO:28)		920	CCA/TCA/TCT/ACA/CGC/TCC/GAG/ ATG/CTG/AGA/TGC/G
E8/4 (SEQ ID NO:29)	rev	920	CGC/ATC/TCA/GCA/TCT/CGG/AGC/ GTG/TAG/ATG/ATG/G

